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Claim Amendments

A listing of all claims follows, with current amendments to claim 1, and the cancellation of claim 2 indicated. The amendment of claim 1 and cancellation of claim 2 are made without prejudice or disclaimer.

1 (currently amended): A method of screening a candidate antiviral agent for antiviral activity comprising

- (a) preparing a first cell culture comprising cells containing a first subgenomic viral replication system, and a second cell culture comprising cells containing a second subgenomic viral replication system; then
 - (b) adding the candidate antiviral agent to each cell culture; then
- (c) incubating the cell cultures under conditions and for a time sufficient to detect an antiviral effect by the candidate antiviral agent on the subgenomic viral replication systems; and
- (d) determining the effect of the candidate antiviral agent on each viral replication system,

wherein the first subgenomic viral replication system is genetically distinct from the second subgenomic viral replication system, and wherein the first and second cell cultures are combined before step (b).

2 (cancelled)

(previously amended): The method of claim 1, further comprising, in step (a), preparing a third cell culture comprising cells not containing a subgenomic viral replication system.

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4 (original): The method of claim 1, wherein at least one of the subgenomic viral replication systems is a replicon.

(original): The method of claim 1, wherein at least one of the subgenomic viral replication systems is a defective genome.

& (original): The method of claim 1, wherein the cells in the first and second cell cultures are mammalian cells and the first and second subgenomic viral replication systems are from mammalian viruses.

ኧ(original): The method of claim &, wherein the mammalian cells are human cells and the mammalian viruses are human viruses.

& (previously amended): The method of claim & wherein the human viruses are selected from the group consisting of hepatitis C virus, yellow fever virus, respiratory syncytial virus, Sindbis virus, poliovirus, Japanese encephalitis virus, hepatitis B virus, human papilloma virus, herpes simplex virus type 1, Epstein-Barr virus, adeno-associated virus, Venezuela encephalitis virus, rubella, coxsackievirus, enterovirus, hepatitis A virus, Dengue fever virus, West Nile virus, tick-borne encephalitis virus, astrovirus, rabies virus, influenza virus A, influenza virus B, measles, mumps, Ebola virus, Marburg virus, La Crosse virus, California encephalitis virus, Hantaan virus, Crimean-Congo virus, Rift Valley fever, Lassa fever, Argentine hemorrhagic fever virus, Bolivian hemorrhagic fever virus, Colorado tick fever, JC virus, BK virus, herpes simplex virus type two, human cytomegalovirus, varicella-zoster virus, human herpes simplex virus type six, human herpes virus type seven, human herpes virus type eight, human adenovirus, HIV-1, HIV-2, HTLV-1, HTLV-2, and human parvovirus.

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g (previously amended): The method of claim χ , wherein the human viruses are selected from the group consisting of hepatitis C virus, yellow fever virus, respiratory syncytial virus, Sindbis virus, poliovirus, Japanese encephalitis virus, hepatitis B virus, human papilloma virus, herpes simplex virus type 1, Epstein-Barr virus, and adenoassociated virus.

10 (previously amended): The method of claim χ , wherein the human viruses are selected from the group consisting of hepatitis C virus, respiratory syncytial virus, yellow fever virus and Sindbis virus.

11-25 (canceled)

10

26 (previously amended): The method of claim 1, further comprising at least a third cell culture comprising cells containing a third subgenomic viral replication system, wherein the third cell culture is also subjected to steps (a), (b), (c) and (d),

wherein each subgenomic viral replication system is genetically distinct from every other subgenomic viral replication system.

11 10 2\(\text{V}\) (original): The method of claim 26, wherein all cell cultures comprising a subgenomic viral replication system are combined before step (b).

28-54 (canceled)

12

\$5 (original): A mixed cell culture comprising a first cell culture comprising cells containing a first subgenomic viral replication system and a second cell culture comprising cells containing a second subgenomic viral replication system.

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56 (original): The mixed cell culture of claim 55, wherein at least one of the subgenomic viral replication systems is a replicon.

by (original): The mixed cell culture of claim \$5, wherein at least one of the subgenomic viral replication systems is a defective genome.

15-58 (original): The mixed cell culture of claim 55, wherein all of the cells of the mixed cell culture are the same cell line.

50 (original): The mixed cell culture of claim 55, wherein the cells of the mixed cell culture comprise more than one cell line.

60 (original): The mixed cell culture of claim 55, wherein all of the cells in the mixed cell culture are mammalian cells and all of the subgenomic viral replication systems are from mammalian viruses.

17 61 (original): The mixed cell culture of claim 60, wherein the mammalian cells are human cells and the mammalian viruses are human viruses.

62 (previously amended): The mixed cell culture of claim 61, wherein the human viruses are selected from the group consisting of hepatitis C virus, yellow fever virus, respiratory syncytial virus, Sindbis virus, poliovirus, Japanese encephalitis virus, hepatitis B virus, human papilloma virus, herpes simplex virus type 1, Epstein-Barr virus, adenoassociated virus, Venezuela encephalitis virus, rubella, coxsackievirus, enterovirus, hepatitis A virus, Dengue fever virus, West Nile virus, tick-borne encephalitis virus, astrovirus, rabies virus, influenza virus A, influenza virus B, measles, mumps, Ebola

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virus, Marburg virus, La Crosse virus, California encephalitis virus, Hantaan virus, Crimean-Congo virus, Rift Valley fever, Lassa fever, Argentine hemorrhagic fever virus, Bolivian hemorrhagic fever virus, Colorado tick fever, JC virus, BK virus, herpes simplex virus type two, human cytomegalovirus, varicella-zoster virus, human herpes simplex virus type six, human herpes virus type seven, human herpes virus type eight, human adenovirus, HIV-1, HIV-2, HTLV-1, HTLV-2, and human parvovirus.

20 63 (previously amended): The mixed cell culture of claim 64, wherein the human viruses are selected from the group consisting of hepatitis C virus, yellow fever virus, respiratory syncytial virus, Sindbis virus, poliovirus, Japanese encephalitis virus, hepatitis B virus, human papilloma virus, herpes simplex virus type 1, Epstein-Barr virus, and adeno-associated virus.

21 64 (previously amended): The mixed cell culture of claim 64, wherein the human viruses are selected from the group consisting of hepatitis C virus, respiratory syncytial

virus, yellow fever virus and Sindbis virus.

65-67 (canceled)

ندنه 68 (original): The mixed cell culture of claim 55, further comprising a third cell culture comprising cells containing a third subgenomic viral replication system.

92 69 (original): The mixed cell culture of claim 68, wherein the mixed cell culture further comprises a fourth cell culture comprising cells containing a fourth subgenomic viral replication system.